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Dated: 5/30/03

Docket No.: GNCA-P03-007 (PATENT)

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In re Patent Application of: Beach et al

Application No.: 10/055797

Group Art Unit: 1638

Filed: January 22, 2002

Examiner: Not Yet Assigned

For:

METHODS AND COMPOSITIONS FOR

RNA INTERFERENCE

INFORMATION DISCLOSURE STATEMENT (IDS)

Commissioner for Patents P.O. Box 1450 Alexandria, VA 22313

Pursuant to 37 CFR 1.56, the attention of the Patent and Trademark Office is hereby directed to the references listed on the attached PTO/SB/08. It is respectfully requested that the information be expressly considered during the prosecution of this application, and that the references be made of record therein and appear among the "References Cited" on any patent to issue therefrom.

This Information Disclosure Statement is filed before the mailing date of a first Office Action on the merits as far as is known to the undersigned.

A copy of each reference on PTO/SB/08 is attached.

While the information and references disclosed in this Information Disclosure Statement may be "material" pursuant to 37 CFR 1.56, it is not intended to constitute an admission that any patent, publication or other information referred to therein is "prior art" for this invention unless specifically designated as such.

Application No.: 10/055797 Docket No.: GNCA-P03-007

In accordance with 37 CFR 1.97(g), the filing of this Information Disclosure Statement shall not be construed to mean that a search has been made or that no other material information as defined in 37 CFR 1.56(a) exists. Applicants further reserve the right to take appropriate action to establish the patentability of the disclosed invention over the listed documents should one or more of the documents be applied against the claims of the present application.

The Commissioner is hereby authorized to charge any deficiency in the fees filed, asserted to be filed or which should have been filed herewith (or with any paper hereafter filed in this application by this firm) to our Deposit Account No. 18/1945, under Order No. GNCA-P03-007.

Dated: May 30, 2003

Respectfully submitted

Matthe VP Vincent

Registration No.: 86,709

ROPES & GRAY LIP One International Place

Boston, Massachusetts 02110-2624

(617) 951-7000 (617) 951-7050 (Fax)

Attorneys/Agents For Applicant

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Initials* No.1	Number-Kind Code ² (if known)	144 55 10004	of Cited Document	Passages or Relevant Figures Appear
AA	6,326,193	12-04-2001	Liu et al.	

		FOREI	GN PATENT	DOCUMENTS		
Examiner		Foreign Patent Document	Publication Date	Name of Patentee or	Pages, Columns, Lines, Where Relevant	
Initials*			MM-DD-YYYY	Applicant of Cited Document	Passages or Relevant Figures Appear	₽
	AB	WO 01/36646	05-25-2001	Cancer Res. Campaign Tech.		
	AC	WO 01/48183	07-05-2001	Devgen NV		
	AD	WO 01/75164	10-11-2001	Whitehead Inst. Biomed. Res.		
	AE	WO 02/44321	06-06-2002	Max-Planck-Gesellschaft		
	AF	WO 02/059300	08-01-2002	J & J Res. Pty Ltd		
	AG	WO 02/068635	09-06-2002	Novartis		

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¹ Applicant's unique citation designation number (optional). ² See attached Kinds Codes of USPTO Patent Documents at www.uspto.gov or MPEP 901.04. ³ Enter Office that issued the document, by the two-letter code (WIPO Standard ST.3). ⁴ For Japanese patent documents, the indication of the year of the reign of the Emperor must precede the application number of the patent document. ⁵ Kind of document by the appropriate symbols as indicated on the document under WIPO Standard ST. 16 if possible. ⁸ Applicant is to place a check mark here if English language Translation is attached.

		OTHER PRIOR ART – NON PATENT LITERATURE DOCUMENTS	
Examiner Initials	Cite No.1	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc), date, page(s), volume-issue number(s), publisher, city and/or country where published.	T²
	АН	Bass, B.L. Double-Stranded RNA as a Template for Gene Silencing. <i>Cell</i> 101, 235-238 (2000).	
	Al	Baulcombe, D.C. RNA as a target and an initiator of post-transcriptional gene silencing in transgenic plants. <i>Plant Mol. Biol.</i> 32, 79-88 (1996).	
	AJ	Baulcombe, D.C. Gene silencing: RNA makes RNA makes no protein. Curr. Biol. 9, R599-R601 (1999).	
	AK	Bohmert, K. et al. AGO1 defines a novel locus of Arabidopsis controlling leaf development. EMBO J. 17, 170-180 (1998).	
	AL	Bosher, J.M. et al. RNA Interference Can Target Pre-mRNA: Consequences for Gene Expression in a Caenorhabditis elegans Operon. <i>Genetics</i> 153, 1245-1256 (Nov. 1999).	
	AM	Bosher, J.M. & Labouesse, M. RNA interference: genetic wand and genetic watchdog. <i>Nat. Cell Biol.</i> 2, E31-36 (2000).	
	AN	Catalanotto, C. et al. Gene silencing in worms and fungi. Nature 404, 245 (2000).	

Examiner	Date	
Signature	Considered	

PTO/SB/08A (10-01)

Approved for use through 10/31/2002.OMB 0651-0031
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ork Reduction Act of 1995, no persons are required to respond to a collection of information unless it contains a valid OMB cont THATTE Complete if Known Substitute for form 1449A/PTO 10/055797 Application Number INFORMATION DISCLOSURE January 22, 2002 Filing Date STATEMENT BY APPLICANT First Named Inventor David Beach Art Unit 1638 (use as many sheets as necessary) Examiner Name Not Yet Assigned 5 GNCA-P03-007 Sheet 2 of Attorney Docket Number

AO	Cogoni, C. & Macino, G. Gene silencing in Neurospora crassa requires a protein homologous to RNA-dependent RNA polymerase. <i>Nature</i> 399, 166-169 (1999).	
AP	Cogoni, C. & Macino, G. Posttranscriptional Gene Silencing in Neurospora by a RecQ DNA Helicase. <i>Science</i> 286, 2342-2344 (1999).	r
AQ	Connelly, J.C. & Leach, D.R. The sbcC and sbcD genes of Escherichia coli encode a nuclease involved in palindrome inviability and genetic recombination. <i>Genes Cell</i> 1, 285-291 (1996).	
AR	Dalmay, T. et al. An RNA-Dependent RNA Polymerase Gene in Arabidopsis is Required for Posttranscriptional Gene Silencing Mediated by a Transgene but Not by a Virus. <i>Cell</i> 101, 543-553 (2000).	
AS	Di Nocera, P.P. & Dawid, I.B. Transient expression of genes introduced into cultured cells of Drosophila. <i>PNAS</i> 80, 7095-7098 (1983).	
AT	Fagard, M. et al. AG01, QDE-2, and RDE-1 are related proteins required for post-transcriptional gene silencing in plants, quelling in fungi, and RNA interference in animals. <i>PNAS</i> 97, 11650-11654 (10 Oct. 2000).	
AU	Fire, A. RNA-triggered gene silencing. <i>Trends Genet.</i> 15, 358-363 (1999).	
AV	Fire, A. et al. Potent and specific genetic interference by double-stranded RNA in Caenorhabditis elegans. <i>Nature</i> 391, 806-811 (1998).	
AW	Fortier, E. & Belote, J.M. Temperature-Dependent Gene Silencing by an Expressed Inverted Repeat in Drosophila. <i>Genesis</i> 26, 240-244 (2000).	
AX	Gillespie, D.E. & Berg, C.A. homeless is required for RNA localization in Drosophila oogenesis and encodes a new member of the DE-H family of RNA-dependent ATPases. <i>Genes Dev.</i> 9, 2495-2508 (1995).	
AY	Guo, S. & Kemphues, K.J. par-1, a Gene Required for Establishing Polarity in C. elegans Embryos, Encodes a Putative Ser/Thr Kinase that is Asymmetrically Distributed. <i>Cell</i> 81, 611-620 (1995).	
AZ	Hamilton, J.A. & Baulcombe, D.C. A Species of Small Antisense RNA in Posttranscriptional Gene Silencing in Plants. <i>Science</i> 286, 950-952 (1999).	
ВА	Hammond, S.M. et al. An RNA-directed nuclease mediates post-transcriptional gene silencing in Drosophila cells. <i>Nature</i> 404, 293-296 (2000).	
ВВ	Hunter, C. Genetics: A touch of elegance with RNAi. Curr. Biol. 9, R440-R442 (1999).	
BC	Jacobsen, S.E. et al. Disruption of an RNA helicase/RNAse III gene in Arabidopsis causes unregulated cell division in floral meristems. <i>Development</i> 126, 5231-5243 (1999).	
BD	Jones, A.L. et al. De novo methylation and co-suppression induced by a cytoplamically replicating plant RNA virus. <i>EMBO J.</i> 17, 6385-6393 (1998).	

Examiner	Date	
Signature	Considered	

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PTO/SB/08A (10-01)
Approved for use through 10/31/2002.OMB 0651-0031
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BE	Jones, L. et al. RNA-DNA Interactions and DNA Methylation in Post-Transcriptional Gene Silencing. <i>Plant Cell</i> 11, 2291-2301 (Dec. 1999).	
BF	Kalejta, R.F. et al. An Integral Membrane Green Fluorescent Protein Marker, Us9-GFP, is Quantitatively Retained in Cells during Propidium Iodide-Based Cell Cycle Analysis by Flow Cytometry. Exp. Cell. Res. 248, 322-328 (1999).	
BG	Kennerdell, J.R. & Carthew, R.W. Use of dsRNA-Mediated Genetic Interference to Demonstrate that frizzled and frizzled 2 Act in the Wingless Pathway. <i>Cell</i> 95, 1017-1026 (1998).	
ВН	Kennerdell, J.R. & Carthew, R.W. Heritable gene silencing in Drosophila using double-stranded RNA. <i>Nat. Biotechnol.</i> 17, 896-898 (2000).	
ВІ	Ketting, R.F. et al. mut-7 of C. elegans, Required for Transposon Silencing and RNA Interference, Is a Homolog of Werner Syndrome Helicase and RNaseD. <i>Cell</i> 99, 133-141 (1999).	
BJ	Kramer, E.R. et al. Activation of the human anaphase-promoting complex by proteins of the CDC20/Fizzy family. <i>Curr. Biol.</i> 8, 1207-1210 (1998).	
ВК	Lam, G. & Thummel, C.S. Inducible expression of double-stranded RNA directs specific genetic interference in Drosophila. <i>Curr. Biol.</i> 10, 957-963 (2000).	
BL	Lohmann, J.U. et al. Silencing of Developmental Genes in Hydra. <i>Dev. Biol.</i> 214, 211-214 (1999).	
ВМ	Matsuda, S. et al. Molecular cloning and characterization of a novel human gene (HERNA) which encodes a putative RNA-helicase. <i>Biochim. Biophys. Acta</i> 1490, 163-169 (2000).	
BN	Misquitta, L. & Paterson, B.M. Targeted disruption of gene function in Drosophila by RNA interference (RNA-i): A role for nautilus in embryonic somatic muscle formation. <i>PNAS</i> 96, 1451-1456 (Feb. 1999).	
ВО	Montgomery, M.K. et al. RNA as a target of double-stranded RNA-mediated genetic interference in Caenorhabditis elegans. <i>PNAS</i> 95, 15502-15507 (Dec. 1998).	
ВР	Montgomery, M.K. & Fire, A. Double-stranded RNA as a mediator in sequence-specific genetic silencing and co-suppression. <i>Trends Genet.</i> 14, 255-258 (1998).	
BQ	Mourrain, P. et al. Arabidopsis SGS2 and SGS3 Genes are Required for Posttranscriptional Gene Silencing and Natural Virus Resistance. <i>Cell</i> 101, 533-542 (2000).	
BR	Ngo, H. et al. Double-stranded RNA induces mRNA degradation in Trypanosoma brucei. PNAS 95, 14687-14692 (Dec. 1998).	
BS	Ratcliff, F. et al. A Similarity Between Viral Defense and Gene Silencing in Plants. <i>Science</i> 276, 1558-1560 (1997).	

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First Named Inventor David Beach

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Yaf Assigned Substitute for form 1449A/PTO INFORMATION DISCLOSURE STATEMENT BY APPLICANT (use as many sheets as necessary) Sheet

BT Sanchez Alvarado, A. & Newmark, P.A. Double-stranded RNA specifically disrupts gexpression during planarian regeneration. PNAS 96, 5049-5054 (April 1999). BU Schneider, I. Cell lines derived from late embryonic stages of Drosophila melanogas Embryol. Exp. Morpho. 27, 353-365 (1972). BV Sharp, P.A. RNAi and double-strand RNA. Genes Dev. 13, 139-141 (1999). BW Shi, H. et al. Genetic interference in Typanosoma brucei by heritable and inducible distranded RNA. RNA 6, 1069-1076 (2000).	ter. J.
Embryol. Exp. Morpho. 27, 353-365 (1972). BV Sharp, P.A. RNAi and double-strand RNA. Genes Dev. 13, 139-141 (1999). BW Shi, H. et al. Genetic interference in Typanosoma brucei by heritable and inducible d stranded RNA. RNA 6, 1069-1076 (2000).	
BW Shi, H. et al. Genetic interference in Typanosoma brucei by heritable and inducible d stranded RNA. RNA 6, 1069-1076 (2000).	louble-
stranded RNA. RNA 6, 1069-1076 (2000).	louble-
BX Shuttleworth, J. & Colman, A. Antisense oligonucleotide-directed cleavage of mRNA Xenopus oocytes and eggs. <i>EMBO J.</i> 7, 427-434 (1988).	in
BY Sijen, T. & Kooter, J.M. Post-transcriptional gene-silencing: RNAs on the attack or or defense? <i>Bioessays</i> 22, 520-531 (2000).	n the
BZ Smardon, A. et al. EGO-1 is related to RNA-directed RNA polymerase and functions line development and RNA interference in C. elegans. <i>Curr. Biol.</i> 10, 169-178 (2000)	
CA Smith, N.A. et al. Total silencing by intron-spliced hairpin RNAs. Nature 407, 319-32	(2000).
CB Tabara, H. et al. RNAi in C. elegans: Soaking in the Genome Sequence. Science 28 432 (1998).	32, 430-
CC Tabara, H. et al. The rde-1 Gene, RNA Interference, and Transposon Silencing in C. Cell 99, 123-132 (1999).	elegans.
CD Tavernarakis, N. et al. Heritable and inducible genetic interference by double-strande encoded by transgenes. <i>Nat. Genet.</i> 24, 180-183 (2000).	ed RNA
CE Timmons, L. & Fire, A. Specific interference by ingested dsRNA. <i>Nature</i> 395, 854 (1	998).
CF Tuschl, T. et al. Targeted mRNA degradation by double-stranded RNA in vitro. Gen. 13, 3191-3197 (1999).	es Dev.
CG Vaucheret, H. et al. Transgene-induced gene silencing in plants. Plant J. 16, 651-65	9 (1998).
CH Wassenegger, M. & Pelissier, T. A model for RNA-mediated gene silencing in higher Plant Mol. Biol. 37, 349-362 (1998).	plants.
CI Waterhouse, P.M. et al. Virus resistance and gene silencing in plants can be induced simultaneous expression of sense and antisense RNA. <i>PNAS</i> 95, 13959-13964 (No.	
CJ Wianny, F. & Zernicka-Goetz, M. Specific interference with gene function by double-RNA in early mouse development. <i>Nature Cell Biol.</i> 2, 70-75 (2000).	stranded
CK Wolf, D.A. & Jackson, P.K. Cell cycle: Oiling the gears of anaphase. <i>Curr. Biol.</i> 8, R (1998).	636-R639

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**Named Inventor David Beach

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CI	L.	Zamore, P.D. et al. RNAi: Double-Stranded RNA Directs the ATP-Dependent Cleavage of	
		mRNA at 21 to 23 Nucleotide Intervals. Cell 101, 25-33 (2000).	

^{*}EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.

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Signature	Considered

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